# WORLD INTELLECTUAL PROPERTY ORGANIZATION International Bureau



#### INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(51) International Patent Classification 7: C12N 15/31, 15/66, 15/63, 1/21, C07K 14/31

(11) International Publication Number:

WO 00/12678

A3

(43) International Publication Date:

9 March 2000 (09.03.00)

(21) International Application Number:

PCT/US99/19726

(22) International Filing Date:

31 August 1999 (31.08.99)

(30) Priority Data:

60/098,964

1 September 1998 (01.09.98) US

(71) Applicant (for all designated States except US): HUMAN GENOME SCIENCES, INC. [US/US]; 9410 Key West Avenue, Rockville, MD 20850 (US).

(72) Inventors; and

(75) Inventors/Applicants (for US only): BAILEY, Camella, C. [US/US]; 1753 Kilbourne Place NW, Washington, DC 20010 (US). CHOI, Gil, H. [CN/US]; 11429 Potomac Oaks Drive, Rockville, MD 20850 (US).

(74) Agents: HOOVER, Kenley, K. et al.; Human Genome Sciences, Inc., 9410 Key West Avenue, Rockville, MD 20850 (US).

(81) Designated States: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, ARIPO patent (GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG).

Published

With international search report.

(88) Date of publication of the international search report:

15 June 2000 (15.06.00)

(54) Title: STAPHYLOCOCCUS AUREUS GENES AND POLYPEPTIDES

(57) Abstract

The present invention relates to novel genes from S. aureus and the polypeptides they encode. Also provided are vectors, host cells, antibodies and recombinant methods for producing the same. The invention further relates to screening methods for identifying agonists and antagonists of S. aureus polypetide activity. The invention additionally relates to diagnostic methods for detecting Staphylococcus nucleic acids, polypeptides and antibodies in a biological sample. The present invention further relates to novel vaccines for the prevention or attenuation of infection by Staphylococcus.

#### FOR THE PURPOSES OF INFORMATION ONLY

Codes used to identify States party to the PCT on the front pages of pamphlets publishing international applications under the PCT.

AL	Albania	ES	Spain	LS	Lesotho	SI	Slovenia
AM	Armenia	FI	Finland	LT	Lithuania	SK	Slovakia
AT	Austria	FR	France	LU	Luxembourg	SN	Senegal
ΑÜ	Australia	GA	Gabon	LV	Latvia	SZ	Swaziland
AZ	Azerbaijan	GB	United Kingdom	MC	Monaco	TD	Chad
BA	Bosnia and Herzegovina	GE	Georgia	MD	Republic of Moldova	TG	Togo
BB	Barbados	GH	Ghana	MG	Madagascar	TJ	Tajikistan
BE	Belgium	GN	Guinea	MK	The former Yugoslav	TM	Turkmenistan
BF	Burkina Faso	GR	Greece		Republic of Macedonia	TR	Turkey
BG	Bulgaria	HU	Hungary	ML	Mali	TT	Trinidad and Tobago
BJ	Benin	IE	Ireland	MN	Mongolia	UA	Ukraine
BR	Brazil	IL	Israel	MR	Mauritania	UG	Uganda
BY	Belarus	IS	Iceland	MW	Malawi	US	United States of America
CA	Canada	IT	Italy	MX	Mexico	UZ	Uzbekistan
CF	Central African Republic	JP	Japan	NE	Niger	VN	Vict Nam
CG	Congo	KE	Kenya	NL	Netherlands	YU	Yugoslavia
СН	Switzerland	KG	Kyrgyzstan	NO	Norway	ZW	Zimbabwe
CI	Côte d'Ivoire	KP	Democratic People's	NZ	New Zealand		
CM	Cameroon		Republic of Korea	PL	Poland		•
CN	China	KR	Republic of Korea	PТ	Portugal		
Cυ	Cuba	KZ	Kazakstan	RO	Romania		
CZ	Czech Republic	LC	Saint Lucia	RU	Russian Federation		
DE	Germany	LI	Liechtenstein	SD	Sudan		
DK	Denmark	LK	Sri Lanka	SE	Sweden		
EE	Estonia	LR	Liberia	SG	Singapore		
							•

#### INTERNATIONAL SEARCH REPORT

International application No. PCT/US99/19726

A. CLASSIFICATION OF SUBJECT MATTER  IPC(7) :C12N 15/31,15/66, 15/63, 1/21: C07K 14/31  US CL :536/23.7; 435/91.41, 320.1, 252.3, 69.1; 530/350								
According to International Patent Classification (IPC) or to both national classification and IPC								
B. FIELDS SEARCHED								
Minimum documentation scarched (classification system followed by classification symbols)								
U.S. : 536/23.7; 435/91.41, 320.1, 252.3, 69.1; 530/350								
Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched								
Electronic o	data base consulted during the international search (r	name of data base and, where practicable	e, search terms used)					
Genbank, Swissprot, PIR60, SPTREMBL9 search terms: sequences corresponding to SEQ ID NO: 2, 4, 6, 8, 10, 12, 14, 16, 18, and 20								
C. DOCUMENTS CONSIDERED TO BE RELEVANT								
Category*	Citation of document, with indication, where a	ppropriate, of the relevant passages	Relevant to claim No.					
Х	EP 0 843 016 A2 (SMITHKLINE BEE May 1998, page 23 (relevant to instan	1-11						
х	EP 0 811 696 A2 (SMITHKINE BEE December 1997, page 8 (relevant to in	1-11						
x	EP 0 826 774 A2 (SMITHKINE BEE March 1998, Figure 1 (relevant to ins	1-11						
_ <u></u>			·					
Further documents are listed in the continuation of Box C. See patent family annex.								
*A* document defining the general state of the art which is not considered to be of particular relevance  *It later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention								
	B* earlier document published on or after the international filing date "X" document of particular relevance; the							
cite	cument which may throw doubts on priority claim(s) or which is sed to establish the publication date of another citation or other scial reason (as specified)	considered novel or cannot be consider when the document is taken alone  "Y" document of perticular relevance, the						
*0* doc	cument referring to an oral disclosure, use, exhibition or other	"Y" document of perticular, relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art						
"P" doc	cument published prior to the international filing date but later than priority date claimed	*&* document member of the same patent family						
Date of the	actual completion of the international search	Date of mailing of the international search report						
10 FEBRU	JARY 2000	0 6 APR 2000						
Commission Box PCT	nailing address of the ISA/US ner of Patents and Trademarks	Authorized officer James Lee For						
Washington Facsimile N	a, D.C. 20231 o. (703) 305-3230	Telephone No. (703) 308-0196						
		1						

### INTERNATIONAL SEARCH REPORT

International application No. PCT/US99/19726

Box I Observations where certain claims were found unessentials (Constitution of the Constitution of the C						
Box I Observations where certain claims were found unsearchable (Continuation of item I of first sheet)						
This international report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:						
1. Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:						
2. Claims Nos.: because they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically:						
3. Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).						
Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)						
This International Scarching Authority found multiple inventions in this international application, as follows:						
Please See Extra Sheet.						
1 As all required additional access for the state of the						
1. As all required additional search fees were timely paid by the applicant, this international search report covers all searchable claims.						
2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite paymen of any additional fee.						
As only some of the required additional search fees were timely paid by the applicant, this international search report covers only those claims for which fees were paid, specifically claims Nos.:						
4. X No required additional search fees were timely paid by the applicant. Consequently, this international search report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:  1-11						
Remark on Protest The additional search fees were accompanied by the applicant's protest.						
No protest accompanied the payment of additional search fees.						

#### INTERNATIONAL SEARCH REPORT

International application No. PCT/US99/19726

## BOX II. OBSERVATIONS WHERE UNITY OF INVENTION WAS LACKING This ISA found multiple inventions as follows:

This application contains the following inventions or groups of inventions which are not so linked as to form a single inventive concept under PCT Rule 13.1. In order for all inventions to be searched, the appropriate additional search fees must be paid.

Group I, claim(s)1-11, drawn to Staphylococcal nucleic acids, proteins encoded by the nucleic acids, vectors comprising the nucleic acids, methods of making the vectors, cells comprising the vectors, and methods of expressing the nucleic acids in transformed cells.

Group II, claim(s) 12 and 13, drawn to an antibody and a cell producing an antibody, both antibodies being specific for the protein encoded by the nucleic acids of Group I.

Group III, claim(s) 16, drawn to a vaccine comprising a protein encoded by the nucleic acids of Group I.

Group IV, claim(s)17, drawn to a method of preventing an infection by administration of a protein encoded by the nucleic acids of Group I.

Group V, claim(s) 18, drawn to a Staphylococcal nucleic acid assay using the nucleic acids of Group I as probes. Group VI, claim(s) 19, drawn to an assay of antibodies specific for Staphylococcal proteins using proteins encoded by the nucleic acids of Group I.

Group VII, claim 20, drawn to an assay of the Staphylococcal proteins of Group I using antibodies specific for the proteins of Group I.

The inventions listed as Groups I-VII do not relate to a single inventive concept under PCT Rule 13.1 because, under PCT Rule 13.2, they lack the same or corresponding special technical features for the following reasons: Groups II-VII constitute the third product, and the second through fifth methods of use of the products of Group I. PCT Rule 13.1 and Annex B do not show that unity of invention exists between a first and second product or method of use.

This application contains claims directed to more than one species of the generic invention. These species are deemed to lack Unity of Invention because they are not so linked as to form a single inventive concept under PCT Rule 13.1. In order for more than one species to be searched, the appropriate additional search fees must be paid. The species are as follows:

The amino acid sequences of the polypeptides shown in Table 1.

The claims are deemed to correspond to the species listed above in the following manner:

All claims of each Group discussed above are drawn to the species indicated above.

The following claims are generic: Claims 1-20

The species listed above do not relate to a single inventive concept under PCT Rule 13.1 because, under PCT Rule 13.2, the species lack the same or corresponding special technical features for the following reasons: Each species is drawn to a different amino acid sequence, and to nucleic acids encoding different amino acid sequences. There is no disclosed relationship between the sequences of each individual species disclosed in Table 1.

Restriction to a single species has been waived sua sponte and the Applicants are permitted to have ten species examined without payment of additional fees. The Applicant's representative Kenley Hoover elected telephonically on 1/27/00 to have the sequences corresponding to SEQ ID NOS: 2, 4, 6, 8, 10, 12, 14, 16, 18, and 20 examined.